

AMENDMENT
US APPLN. NO. 10/730,172

Amendments to the specification:

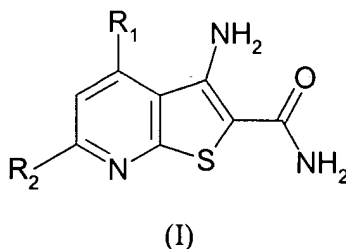
Please replace the paragraph at page 1, line 5 with the following rewritten paragraph:

--This application is a continuation in part of US nonprovisional application No 10,453,175 filed June 3, 2003 under 35 U.S.C 120 which claims benefit to US provisional application No. 60/386,312 filed June 6, 2002.--

Amendments to the Claims:

Listing of Claims as amended:

24. (New) A compound according to general formula (I):



wherein:

R₁ is

- (a) phenyl or heteroaryl selected from furanyl, thienyl, pyridyl, pyrrolyl, imidazolyl and benzofuranyl, optionally substituted with one to two R₃,
- (b) heterocyclyl selected from 1-piperidinyl, 1-piperazinyl, 1-pyrrolidinyl and 4-morpholinyl, optionally substituted with one to two groups selected from C₁₋₆alkyl, -CO₂C₁₋₅alkyl, phenyl, benzyl, -OH and -C(O)heteroaryl, wherein the heteroaryl is selected from furanyl, thienyl, pyridyl and pyrrolyl,
- (c) R₆(CH₂)_mO-,
- (d) R₆OCH₂-,
- (e) R₆(CH₂)_mNH-,

- (f) $R_6(CH_2)_p(CH=CH)_m-$ or pyridyl $(CH_2)_p(CH=CH)_m-$
- (g) C_{1-6} alkyl, optionally partially or fully halogenated and optionally substituted with one to two R_9 ,
- (h) C_{3-6} cycloalkyl
- (i) C_{1-8} alkoxy, optionally partially or fully halogenated and optionally substituted with one to two R_9 ,
- (j) C_{1-8} alkylS(O) $_n-$, optionally partially or fully halogenated and optionally substituted with one to two R_9 ,
- (k) $-N(R_4)(R_5)$, or
- (l) $-C(O)NHR'$, wherein R' is R_6 , pyridyl;

R_2 is

- (a) heterocyclyl $(CH_2)_m-$ wherein said heterocycle is selected from piperidinyl, piperazinyl, morpholinyl, azepanyl, pyrrolidinyl, 1,4-diazacycloheptanyl, azepanyl, 2,5-diazabicyclo[2.2.1]heptanyl, oxazepanyl and thiomorpholino and is optionally substituted with one to three R_7 ,
- (b) heterocyclyl CH_2O- wherein the heterocyclyl is selected from 1-piperidinyl, 1-piperazinyl, 4-morpholinyl and 1-pyrrolidinyl, optionally substituted with C_{1-6} alkyl;

R_3 is chosen from C_{1-6} alkyl, C_{1-6} alkoxy, hydroxy C_{1-6} alkyl, halogen, $-CN$, $-CO_2H$, $-CO_2C_{1-6}$ alkyl, $-S(O)_nC_{1-6}$ alkyl, $-NO_2$, $-OH$, $-CF_3$, $-N(R_4)(R_5)$, $-NHC(O)NHC_{1-6}$ alkyl, $-C(O)N(R_4)(R_5)$ and phenyl optionally substituted with halogen, C_{1-6} alkyl, $-CN$ or C_{1-6} alkoxy;

R_4 and R_5 are independently selected from H, C_{1-6} alkyl, $-C_{0-3}$ alkyl C_{3-6} cycloalkyl, $-C_{0-3}$ alkylheteroaryl selected from the list consisting of benzothiophenyl, furanyl, tetrazolyl, pyridyl, $-C_{0-3}$ alkylheterocyclyl selected from the list consisting of piperidinyl and morpholinyl, $-C_{0-3}$ alkylphenyl(R_6), 2-methylcyclohexyl, $-C(O)C_{1-6}$ alkyl, $-SO_2C_{1-6}$ alkyl, phenyl, pyridyl, piperidinyl, phenylethyl optionally substituted with hydroxymethyl, $(CH_3)_3COC(O)-$, $-CH_2CO_2Me$, $-C_{1-6}$ alkylOH, $-C_{1-6}$ alkylNMe $_2$, or alternatively R_4 and R_5

with the atom to which they are attached can be fused together to form a heterocyclic ring which may be substituted with an OH group;

R_6 is a phenyl group optionally substituted with one to three groups selected from halogen, C_{1-6} alkyl, -CN, $-CO_2C_{1-6}$ alkyl, $-CO_2H$, $-C(O)NR_4R_5$, $-CH_2N(R_4)(R_5)$, $-SO_2N(R_4)(R_5)$, $-NHSO_2C_{1-6}$ alkyl, $-SO_2C_{1-6}$ alkyl, $-NO_2$, $-OH$, $-NH_2$, $-CF_3$, OCF_2 , OCF_3 , $OBenzyl$, C_{1-6} alkoxy, a heteroaryl group selected from the list consisting of pyridyl, pyrazine, imidazolyl and thiazolyl which may be further substituted by an R_4 group, phenyl, a heterocyclic group, or alternatively when R_6 is phenyl two of its adjacent carbon atoms may be bridged by an $-OCH_2O-$ or an $-OCF_2O-$ group, or R_6 is C_{3-6} cycloalkyl, $-CH_2OH$, naphthalene-2-yl, naphthalene-1-yl or 2-thienyl;

R_7 is $R_6CH(OH)CH_2NH-$, C_{1-6} alkyl

R_9 is selected from oxo, $-OH$, $-NR_4R_5$, $-CO_2H$ and C_{1-6} alkoxy;

m is 0 or 1;

n is 0, 1 or 2; and

p is 0, 1, 2 or 3

or pharmaceutically acceptable salts pharmaceutically acceptable salts, isomers or tautomers thereof.

25. The compounds of general formula I of claim 24 wherein:

R_1 is

- (a) phenyl or heteroaryl selected from furanyl, thienyl, pyridyl, pyrrolyl, imidazolyl and benzofuranyl, optionally substituted with one to two R_3 ,
- (b) heterocyclyl selected from 1-piperidinyl, 1-piperazinyl, 1-pyrrolidinyl and 4-morpholinyl, optionally substituted with one to two groups selected from C_{1-6} alkyl, $-CO_2C_{1-5}$ alkyl, phenyl, benzyl, $-OH$ and $-C(O)$ heteroaryl, wherein the heteroaryl is selected from furanyl, thienyl, pyridyl and pyrrolyl,
- (c) $R_6(CH_2)_mO-$,
- (d) R_6OCH_2- ,
- (e) $R_6(CH_2)_mNH-$,
- (f) $R_6(CH_2)_p(CH=CH)_m-$ or pyridyl $(CH_2)_p(CH=CH)_m-$
- (g) C_{1-6} alkyl, optionally partially or fully halogenated and optionally substituted with one to two R_9 ,
- (h) C_{3-6} cycloalkyl
- (i) C_{1-8} alkoxy, optionally partially or fully halogenated and optionally substituted with one to two R_9 ,
- (j) C_{1-8} alkyl $S(O)_n-$, optionally partially or fully halogenated and optionally substituted with one to two R_9 ,
- (k) $-N(R_4)(R_5)$, or
- (l) $-C(O)NHR'$, wherein R' is R_6 , pyridyl or $-CH_3$;

R_2 is piperidinyl $(CH_2)_m-$ optionally substituted with one to three R_7 ,

R_3 is chosen from C_{1-6} alkyl, C_{1-6} alkoxy, hydroxy C_{1-6} alkyl, halogen, $-CN$, $-CO_2H$, $-CO_2C_{1-6}$ alkyl, $-S(O)_nC_{1-6}$ alkyl, $-NO_2$, $-OH$, $-CF_3$, $-N(R_4)(R_5)$, $-NHC(O)NHC_{1-6}$ alkyl, $-C(O)N(R_4)(R_5)$ and phenyl optionally substituted with halogen, C_{1-6} alkyl, $-CN$ or C_{1-6} alkoxy;

R_4 and R_5 are independently selected from, $-C_{0-3}$ alkylheteroaryl, $-C_{0-3}$ alkylheterocyclyl, $-C_{0-3}$ alkylphenyl(R_6), -2 -methylcyclohexyl, $-C(O)C_{1-6}$ alkyl, $-SO_2C_{1-6}$ alkyl, phenyl, pyridyl, piperidinyl, phenylethyl optionally substituted with hydroxymethyl, $(CH_3)_3COC(O)-$, $-CH_2CO_2Me$, $-C_{1-6}$ alkyl OH , $-C_{1-6}$ alkyl NMe_2 , or alternatively R_4 and R_5 with the atom to which they are attached can be fused together to form a heterocyclic ring which may be

substituted with an OH group, provided that R₄ or R₅ are not methyl or benzyl and R₄ and R₅ are not both H;

R₆ is a phenyl group substituted with one to three groups selected from C₁₋₆ alkyl -CN, -CO₂H, -CH₂N(R₄)(R₅), -SO₂N(R₄)(R₅), -C(O)NR₄R₅, NHSO₂C₁₋₆alkyl, -SO₂C₁₋₆alkyl, -CF₃, OCF₂, OCF₃, OBenzyl, C₂₋₆alkoxy, a heteroaryl group selected from the list consisting of pyridyl, pyrazinyl, imidazolyl and thiazolyl which may be further substituted by an R₄ group, phenyl, a heterocyclic group, or alternatively when R₆ is phenyl two of its adjacent carbon atoms may be bridged by an -OCH₂O- or an -OCF₂O- group;

R₇ is R₆CH(OH)CH₂NH-, C₁₋₆alkyl

R₉ is selected from oxo, -OH, -NR₄R₅, -CO₂H and C₁₋₆alkoxy;

m is 0 or 1;

n is 0, 1 or 2; and

p is 0, 1, 2 or 3.

26. The compounds of general formula I of claim 24 wherein:

R₁ is C₁₋₆alkyl, optionally partially or fully halogenated and optionally substituted with one to two R₉;

R₂ is piperdiny[(CH₂)_m] - optionally substituted with one to three R₇;

R₃ is chosen from C₁₋₆alkyl, C₁₋₆alkoxy, hydroxyC₁₋₆alkyl, halogen, -CN, -CO₂H, -CO₂C₁₋₆alkyl, -S(O)_nC₁₋₆alkyl, -NO₂, -OH, -CF₃, -N(R₄)(R₅), -NHC(O)NHC₁₋₆alkyl, -C(O)N(R₄)(R₅) and phenyl optionally substituted with halogen, C₁₋₆alkyl, -CN or C₁₋₆alkoxy;

R₄ and R₅ are independently selected from, -C₀₋₃alkylheteroaryl, -C₀₋₃alkylheterocyclyl, -C₀₋

alkylphenyl(R₆), -2-methylcyclohexyl, -C(O)C₁₋₆alkyl, -SO₂C₁₋₆alkyl, phenyl, pyridyl, piperidinyl, phenylethyl optionally substituted with hydroxymethyl, (CH₃)₃COC(O)-, -CH₂CO₂Me, -C₁₋₆alkylOH, -C₁₋₆alkylNMe₂, or alternatively R₄ and R₅ with the atom to which they are attached can be fused together to form a heterocyclic ring which may be substituted with an OH group, provided that R₄ or R₅ are not methyl or benzyl and R₄ and R₅ are not both H;

R₆ is a phenyl group substituted with one to three groups selected from C₁₋₆ alkyl -CN, -CO₂H, -CH₂N(R₄)(R₅), -SO₂N(R₄)(R₅), -C(O)NR₄R₅, NHSO₂C₁₋₆alkyl, -SO₂C₁₋₆alkyl, -CF₃, OCF₂, OCF₃, OBenzyl, C₂₋₆alkoxy, a heteroaryl group selected from the list consisting of pyridyl, pyrazinyl, imidazolyl and thiazolyl which may be further substituted by an R₄ group, phenyl, a heterocyclic group, or alternatively when R₆ is phenyl two of its adjacent carbon atoms may be bridged by an -OCH₂O- or an -OCF₂O- group;

R₇ is R₆CH(OH)CH₂NH-, C₁₋₆alkyl

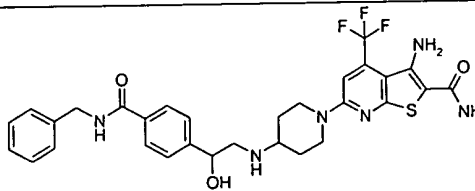
R₉ is selected from oxo, -OH, -NR₄R₅, -CO₂H and C₁₋₆alkoxy;

m is 0 or 1;

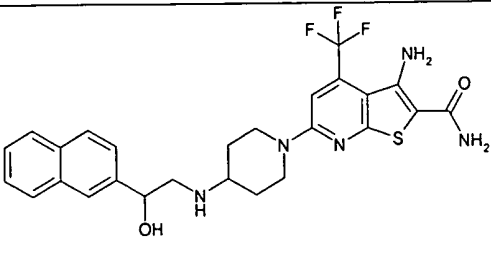
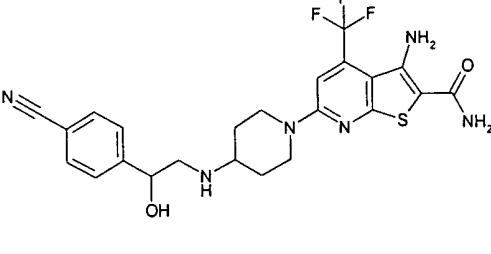
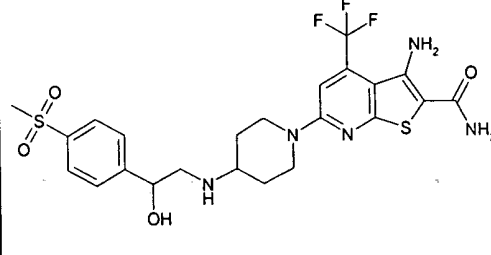
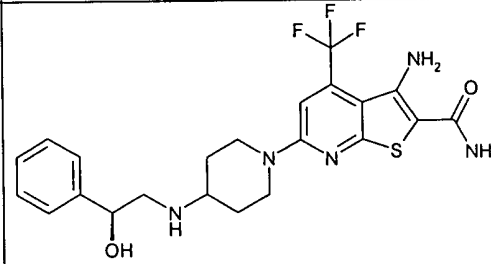
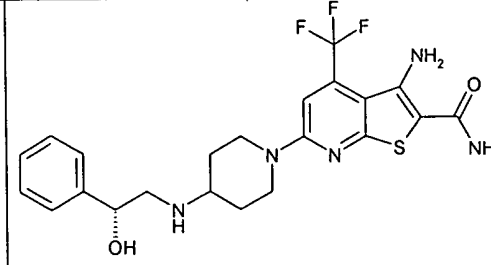
n is 0, 1 or 2; and

p is 0, 1, 2 or 3.

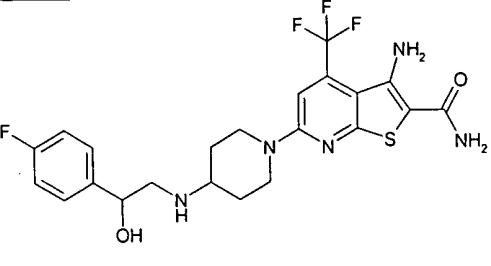
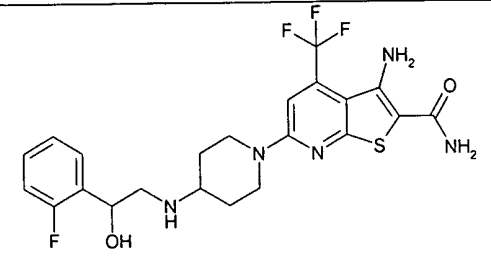
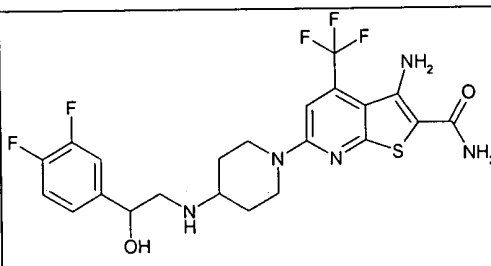
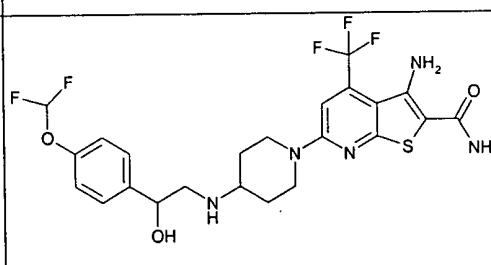
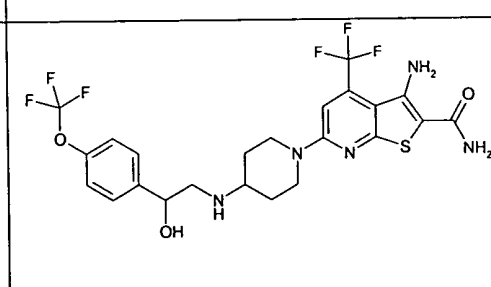
27. A compound selected from the list consisting of:

<p>3-Amino-6-{4-[2-(4-benzylcarbamoyl-piperidin-1-yl)-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
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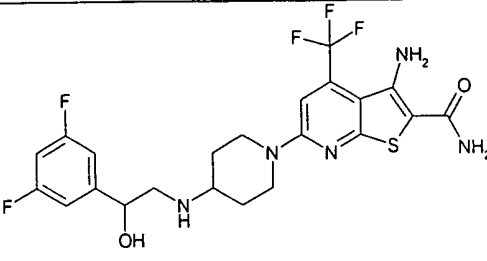
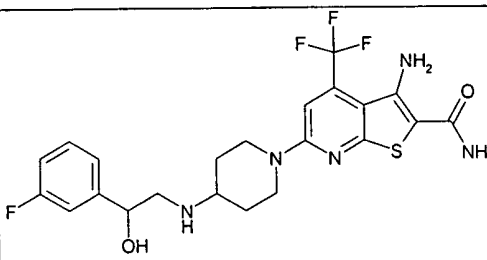
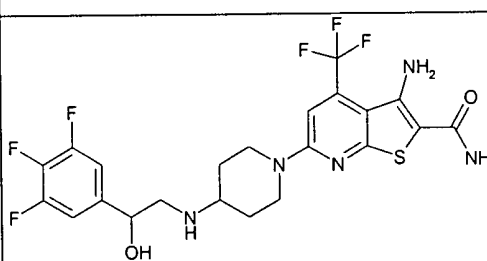
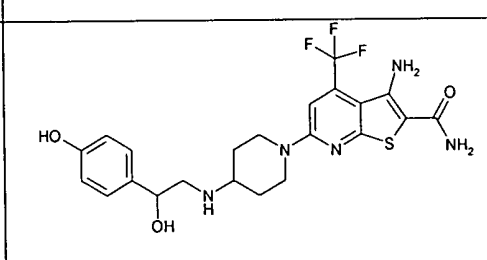
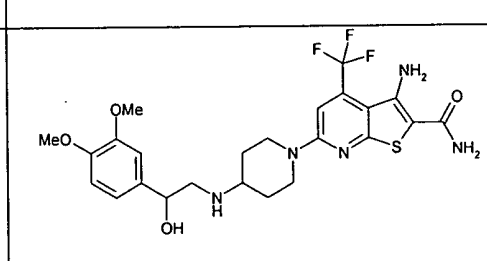
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<p>3-Amino-6-[4-(2-hydroxy-2-naphthalen-2-yl-ethylamino)-piperidin-1-yl]-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-(4-cyano-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-(4-methanesulfonyl-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-[4-((S)-2-hydroxy-2-phenyl-ethylamino)-piperidin-1-yl]-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-[4-((R)-2-hydroxy-2-phenyl-ethylamino)-piperidin-1-yl]-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	

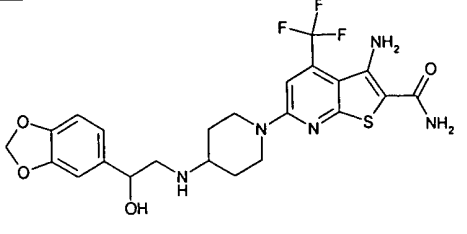
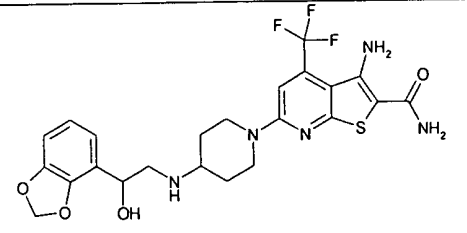
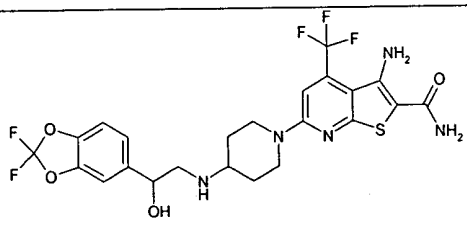
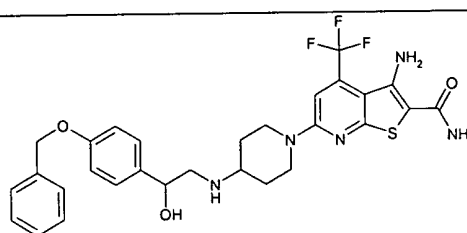
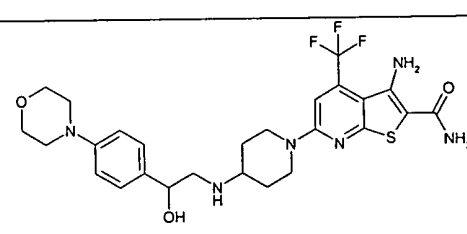
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	3-Amino-6-{4-[2-(4-fluoro-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3- <i>b</i>]pyridine-2-carboxylic acid amide	
	3-Amino-6-{4-[2-(2-fluoro-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3- <i>b</i>]pyridine-2-carboxylic acid amide	
	3-Amino-6-{4-[2-(3,4-difluoro-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3- <i>b</i>]pyridine-2-carboxylic acid amide	
	3-Amino-6-{4-[2-(4-difluoromethoxy-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3- <i>b</i>]pyridine-2-carboxylic acid amide	
	3-Amino-6-{4-[2-hydroxy-2-(4-trifluoromethoxy-phenyl)-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3- <i>b</i>]pyridine-2-carboxylic acid amide	

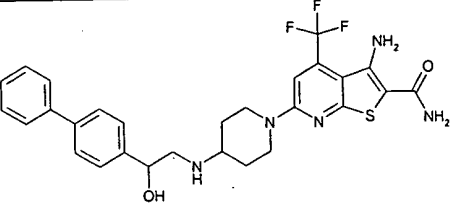
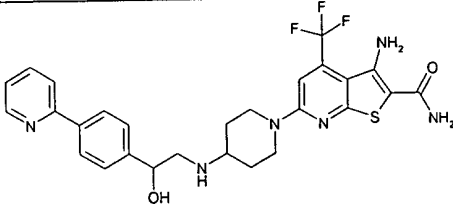
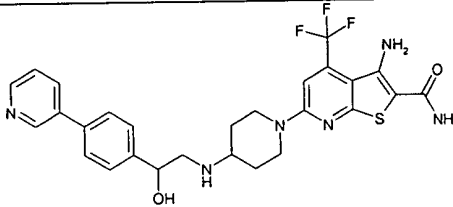
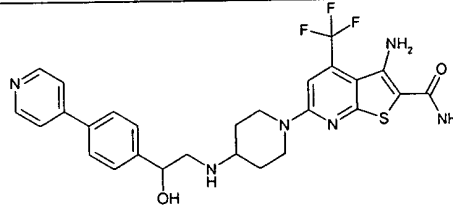
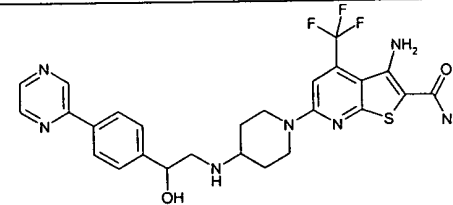
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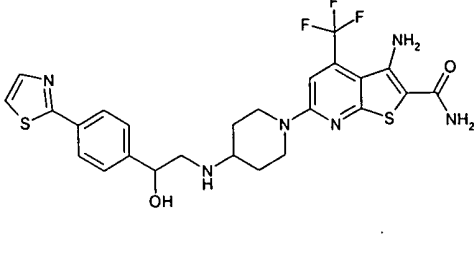
	3-Amino-6-{4-[2-(3,5-difluoro-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3- <i>b</i>]pyridine-2-carboxylic acid amide	
	3-Amino-6-{4-[2-(3-fluoro-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3- <i>b</i>]pyridine-2-carboxylic acid amide	
	3-Amino-6-{4-[2-hydroxy-2-(3,4,5-trifluoro-phenyl)-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3- <i>b</i>]pyridine-2-carboxylic acid amide	
	3-Amino-6-{4-[2-hydroxy-2-(4-hydroxy-phenyl)-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3- <i>b</i>]pyridine-2-carboxylic acid amide	
	3-Amino-6-{4-[2-(3,4-dimethoxy-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3- <i>b</i>]pyridine-2-carboxylic acid amide	

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<p>3-Amino-6-[4-(2-benzo[1,3]dioxol-5-yl-2-hydroxy-ethylamino)-piperidin-1-yl]-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-[4-(2-benzo[1,3]dioxol-4-yl-2-hydroxy-ethylamino)-piperidin-1-yl]-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-(2,2-difluoro-benzo[1,3]dioxol-5-yl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-(4-benzyloxy-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-hydroxy-2-(4-morpholin-4-yl-phenyl)-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	

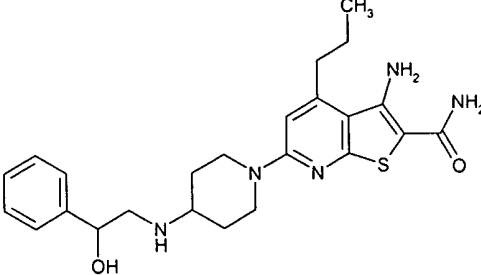
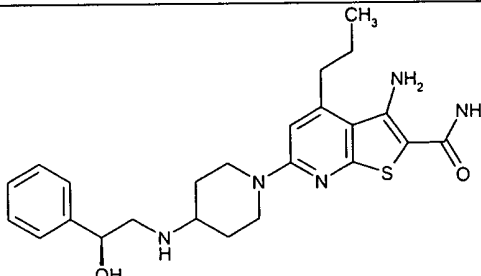
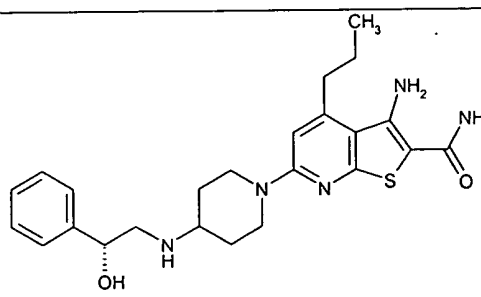
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<p>3-Amino-6-[4-(2-biphenyl-4-yl-2-hydroxy-ethylamino)-piperidin-1-yl]-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-hydroxy-2-(4-pyridin-2-yl-phenyl)-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-hydroxy-2-(4-pyridin-3-yl-phenyl)-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-hydroxy-2-(4-pyridin-4-yl-phenyl)-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-hydroxy-2-(4-pyrazin-2-yl-phenyl)-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	

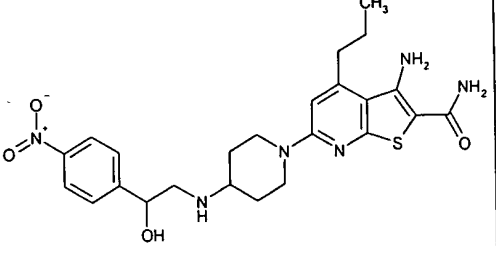
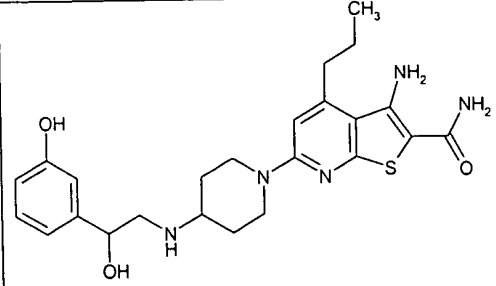
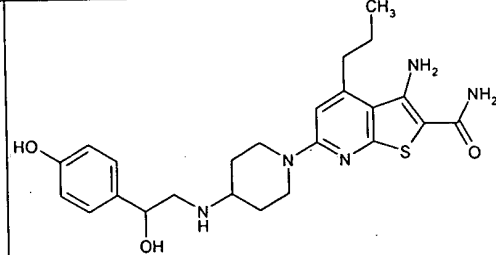
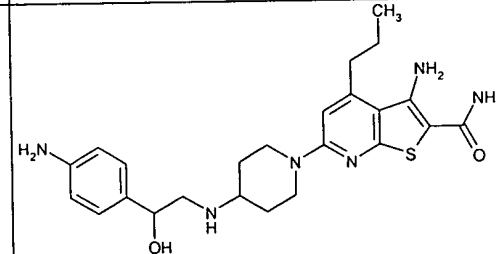
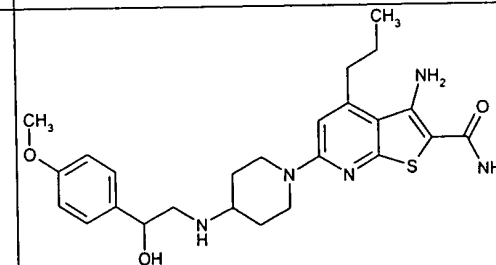
<p>3-Amino-6-{4-[2-hydroxy-2-(4-thiazol-2-yl-phenyl)-ethylamino]-piperidin-1-yl}-4-trifluoromethyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
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or pharmaceutically acceptable salts, tautomers and isomers thereof.

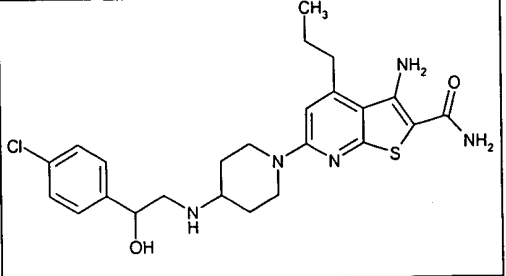
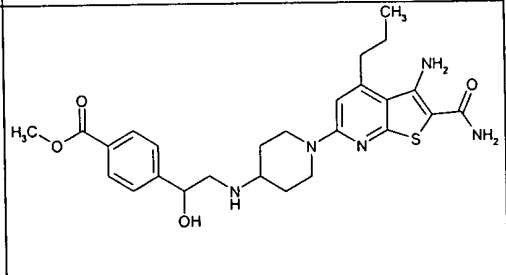
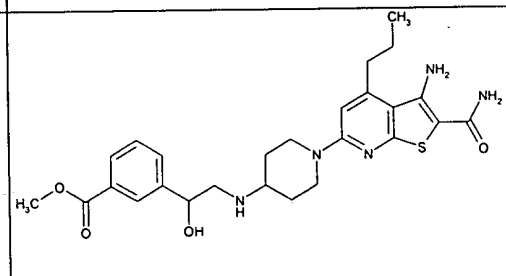
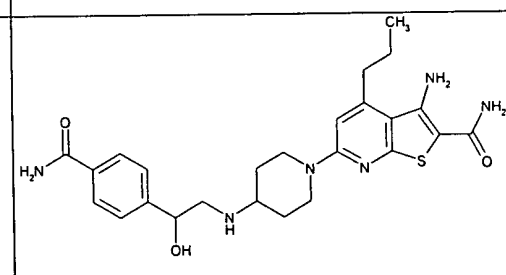
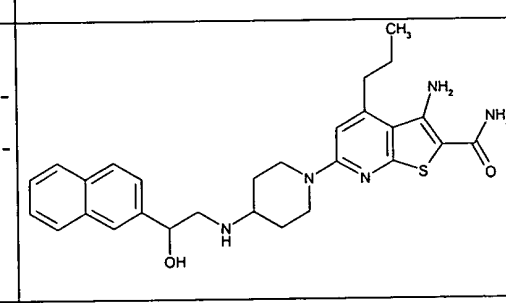
28. A compound selected from the list consisting of:

<p>3-Amino-6-[4-(2-hydroxy-2-phenyl-ethylamino)-piperidin-1-yl]-4-propyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-[4-((S)-2-hydroxy-2-phenyl-ethylamino)-piperidin-1-yl]-4-propyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-[4-((R)-2-hydroxy-2-phenyl-ethylamino)-piperidin-1-yl]-4-propyl-thieno[2,3-<i>b</i>]pyridine-2-carboxylic acid amide</p>	

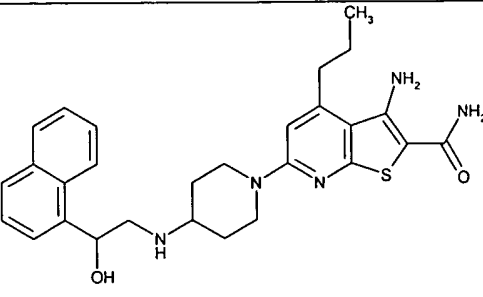
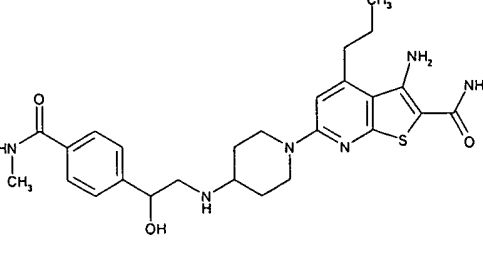
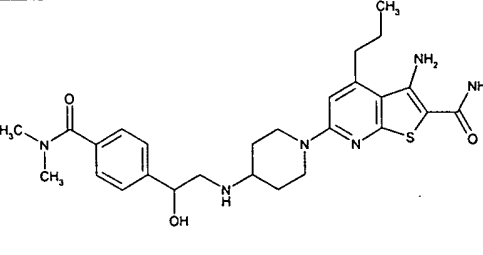
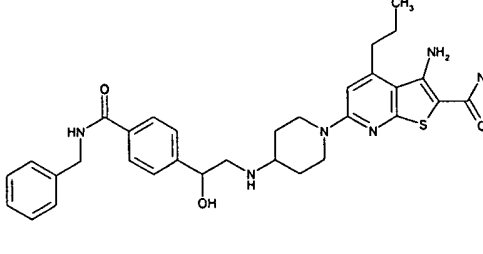
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<p>3-Amino-6-{4-[2-hydroxy-2-(4-nitro-phenyl)-ethylamino]-piperidin-1-yl}-4-propyl-thieno[2,3-b]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-hydroxy-2-(3-hydroxy-phenyl)-ethylamino]-piperidin-1-yl}-4-propyl-thieno[2,3-b]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-hydroxy-2-(4-hydroxy-phenyl)-ethylamino]-piperidin-1-yl}-4-propyl-thieno[2,3-b]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-(4-amino-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-propyl-thieno[2,3-b]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-hydroxy-2-(4-methoxy-phenyl)-ethylamino]-piperidin-1-yl}-4-propyl-thieno[2,3-b]pyridine-2-carboxylic acid amide</p>	

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<p>3-Amino-6-{4-[2-(4-chloro-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-propyl-thieno[2,3-b]pyridine-2-carboxylic acid amide</p>	
<p>4-{2-[1-(3-Amino-2-carbamoyl-4-propyl-thieno[2,3-b]pyridin-6-yl)-piperidin-4-ylamino]-1-hydroxy-ethyl}-benzoic acid methyl ester</p>	
<p>3-{2-[1-(3-Amino-2-carbamoyl-4-propyl-thieno[2,3-b]pyridin-6-yl)-piperidin-4-ylamino]-1-hydroxy-ethyl}-benzoic acid methyl ester</p>	
<p>3-Amino-6-{4-[2-(4-carbamoyl-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-propyl-thieno[2,3-b]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-[4-(2-hydroxy-2-naphthalen-2-yl-ethylamino)-piperidin-1-yl]-4-propyl-thieno[2,3-b]pyridine-2-carboxylic acid amide</p>	

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<p>3-Amino-6-[4-(2-hydroxy-2-naphthalen-1-yl-ethylamino)-piperidin-1-yl]-4-propyl-thieno[2,3-b]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-hydroxy-2-(4-methylcarbamoyl-phenyl)-ethylamino]-piperidin-1-yl}-4-propyl-thieno[2,3-b]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-(4-dimethylcarbamoyl-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-propyl-thieno[2,3-b]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-(4-benzylcarbamoyl-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-propyl-thieno[2,3-b]pyridine-2-carboxylic acid amide</p>	
<p>3-Amino-6-{4-[2-(3-carbamoyl-phenyl)-2-hydroxy-ethylamino]-piperidin-1-yl}-4-propyl-thieno[2,3-b]pyridine-2-carboxylic acid amide</p>	